## ABSTRACT

The invention relates to a method of modulating neutralizing antibodies formation against an heterologous protein. The invention also relates to the use of such method to induce a tolerisation of the immune system towards said protein, such tolerisation being useful to allow long-term gene therapy or transgene expression. The invention also relates to the use of a method of the invention to provide an animal with a reproducible functional inactivation phenotype of an endogenous protein of the animal.